Serial No. Unknown

Int'l Application No. PCT/GB2003/003245

Filing Date Filed Herewith
Int'l Filing Date July 21, 2003
Examiner Unknown
Attorney Docket No. 142.018US01

Title: METHOD AND APPARATUS FOR INVESTIGATING HISTOLOGY OF EPITHELIAL TISSUE Page 3 of 8

Amendments to the Claims:

This listing of claims will replace all prior versions and listings of claims in the application:

Listing of Claims:

1. (Original) A method for monitoring the presence of selected chromophores in a sample of epithelial tissue, independent of the amount of a predetermined chromophore, the method comprising:

illuminating an area of tissue by projecting light from a light source of at least two different wavelengths λ_1 , λ_2 ;

receiving light remitted by the illuminated area of tissue at a photoreceptor; analysing the received light to obtain a measurement $R_i(\lambda)$ for each wavelength and then calculating:

 $Z = \frac{R_i(\lambda_1)}{R_i(\lambda_2)^l}$ where *l* is chosen such that Z is independent of the amount of predetermined chromophore.

- 2. (Original) A method according to claim 1, in which $R_i(\lambda)$ is calculated by analysing the received light to identify and measure the proportion of light of each wavelength remitted from the tissue $I_r(\lambda)$; and calculating the ratio of light at each wavelength returned from the tissue $R_i(\lambda)$.
- 3. (Currently amended) A method according to claim 1 or $\frac{1}{2}$, in which l is calculated

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such that

$$Z = \frac{R_i (c, h, \lambda_1)^j}{R_i (c, h, \lambda_2)^{jk}} = \frac{R_i (\lambda_1)^j}{R_i (\lambda_2)^{jk}} = \frac{R_i (\lambda_1)^j}{R_i (\lambda_2)^l}$$
 where j and k are such that:

 $2j\alpha(\lambda_1) = 2kj\alpha(\lambda_2) = 1$ where $\alpha(\lambda_1)$ and $\alpha(\lambda_2)$ are the absorbtion coefficients for the predetermined chromophore at each wavelength.

- 4. (Currently amended) A method according to any one of the preceding claims claim 1, in which the predetermined chromophore is melanin.
- 5. (Currently amended) A method according to any one of claims 1 to 4 claim 1, in which the predetermined chromophore is haemoglobin.
- 6. (Currently amended) A method according to any one of the preceding claims claim 1, in which the epithelial tissue is skin.
- 7. (Currently amended) A method according to any one of the preceding claims claim 1, in which the wavelengths λ_1 , λ_2 are chosen such that a change in collagen level causes a relatively small change in the absorbtion of λ_1 , and a relatively large change in the absorbtion of λ_2 .
- 8. (Original) A method according to claim 7, in which the difference between the two

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wavelengths λ_1 , λ_2 is at least 200 nm.

9. (Original) A method according to claim 8, in which the wavelengths are substantially 700 nm and 940nm respectively.

10. (Original) A method of forming an image of an area of epithelial tissue independent of the amount of a predetermined chromophore in the tissue, locations, formed by obtaining Z for a plurality of locations within the area, Z being obtained by illuminating an area of tissue by projecting light from a light source of at least two different wavelengths λ_1 , λ_2 ;

receiving light remitted by the illuminated area of tissue at a photoreceptor; analysing the received light to analysing the received light to obtain a measurement $R_{i}(\lambda)$ for each wavelength and then calculating:

 $Z = \frac{R_l(\lambda_1)}{R_l(\lambda_2)^l}$ where l is chosen such that Z is independent of the amount of predetermined chromophore; and mapping the amounts Z at positions indicative of the location within the area of the measurement.

11. (Original) A method according to claim 10, in which $R_i(\lambda)$ is calculated by analysing the received light to identify and measure the proportion of light of each wavelength remitted from the tissue $I_r(\lambda)$; and calculating the ratio of light at each

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wavelength returned from the tissue $R_{i}(\lambda)$.

12. (Currently amended) A method according to claim 10 or 11, in which l is calculated

such that
$$Z = \frac{R_d(c, h, \lambda_1)^j}{R_d(c, h, \lambda_2)^{jk}} = \frac{R_l(\lambda_1)^j}{R_l(\lambda_2)^{jk}} = \frac{R_l(\lambda_1)^l}{R_l(\lambda_1)^l}$$
 where j and k are such that

 $2j\alpha(\lambda_1) = 2kj\alpha(\lambda_2) = 1$ where $\alpha(\lambda_1)$ and $\alpha(\lambda_2)$ are the absorbtion coefficients for the predetermined chromophore at each wavelength.

13. (Currently amended) A method according to any one of the preceding claims claim

10, in which the at least two sets of calculations $Z = \frac{R_i(\lambda_1)}{R_i(\lambda_1)^i}$ are carried out, a first

calculation with l, such that Z is independent of the amount of a first predetermined chromophore, and a second calculation with l_2 such that Z is independent of the amount of a second predetermined chromophore.

14. (Currently amended) A method according to any one of the preceding claims claim 10 in which the light source used to illuminate the tissue, is of at least three wavelengths, $\lambda_1, \lambda_2, \lambda_3$ and at least three pairs of calculations of Z are made,

namely
$$Z = \frac{R_t(\lambda_1)}{R_t(\lambda_2)^{l_1}}$$
, $Z = \frac{R_t(\lambda_2)}{R_t(\lambda_3)^{l_2}}$, $Z = \frac{R_t(\lambda_1)}{R_t(\lambda_3)^{l_3}}$ where $l_1 l_2 l_3$ are each chosen such

that Z is independent of the amount of the predetermined chromophore for the respective pair of wavelengths.

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15. (Original) Apparatus for monitoring the presence of selected chromophores in a sample of epithelial tissue, independent of the amount of a predetermined chromophore comprising a light source for illuminating tissue with light of at least two different wavelengths λ_1 , λ_2 ;

a photoreceptor for receiving images remitted by the illuminated area of tissue at a photoreceptor; and

microprocessor means for analysing the received light to identify and measure the proportion of light of each wavelength remitted from the tissue $I_r(\lambda)$;

calculating the ratio of light at each wavelength returned from the tissue $R_i(\lambda)$, and then calculating: $Z = \frac{R_l(\lambda_1)}{R_l(\lambda_2)^l}$ where *l* is chosen such that Z is independent of the amount of predetermined chromophore.

16. (Original) Apparatus according to claim 15, also comprising image creation means for receiving a plurality of values of Z, each for a specified location on the tissue, and providing a mapped image representing the value of Z at the plurality of locations on the tissue.